Application. No.: 10/629,308

Amendment Dated May 2, 2008

Reply to Office Action of November 5, 2007

Amendments to the Claims: This listing of claims will replace all prior versions, and listings, of claims in the application

Listing of Claims:

 (Currently Amended) A sterile aqueous pharmaceutical composition for parenteral administration of propofol, said composition comprising about 1% (w/v) propofol, and an excipient comprising Poloxamer-188, and one or more additional excipients, in combination with excipients, said excipients including:

7% to 9% (w/v) poloxamer component consisting essentially of Poloxamer 188: 2% to 4% (w/v) polyethylene glycol; and

not more than 1% (w/v) lipid

wherein <u>said exciplents comprise not more than 15% (w/v) of said composition</u> and said composition is stored in a container having a means for dispensing the composition, and wherein the total propofol degradants of the solution when maintained at 25 °C, 40 °C, or 60 °C for 4 weeks are present in an amount of less than 5% (w/v) of said composition.

- 2 5. (Canceled)
- (Currently Amended) The composition of Claim ₹1, wherein÷
- a)-said-composition-comprises poloxamer-188, said polyethylene glycol comprises polyethylene glycol 400, and prepylene glycol:
  - (†)wherein said eemposition-excipients further comprise one or more compounds selected from the group consisting of citric acid, disodium edetate, metabisulfate, benzyl alcohol, propylene glycol, an antioxidant, a preservative, an antimicrobial agent, and a microbicidal.±
  - b) said composition comprises poloxamer 188 and polyethylene gycol 400:
    - (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;

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c) said composition comprises poloxamer 188 (8% w/v), polyethylene glycol 400 (4% w/v), and propylene glycol (1% w/v):

- (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- d) said composition comprises poloxamer 188 (8% w/v), polyethylene glycol 400 (3% w/v), and propylene glycol (1% w/v):
  - (I) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- e) said composition comprises poloxamer 188 (8% w/v), polyethylene glycol 400 (2% w/v), and propylene-glycol (1% w/v):
  - (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol. an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- f) said composition comprises poloxamer 188 (8% w/v) and polyethylene glycol 400 (3% w/v):
  - (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- g) said composition comprises poloxamer 188 (8% w/v) and polyethylene glycol 400 (2% w/v):
  - (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;

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h) said-composition comprises poloxamer 188 (8% w/v) and polyethylene glycol 400 (4% w/v):

- (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- i) said composition comprises poloxamer 188 (7% w/v), polyethylene glycol 400 (3% w/v), and propylene glycol (1% w/v):
  - (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- j) said-composition-comprises polexamer 188 (7% w/v) and polyethylene glycol 400 (3% w/v):
  - (t) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- I) said composition comprises poloxamer 188 (7% w/v), polyethylene glycol 400 (2% w/v), and propylene glycol (1% w/v):
  - (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl-alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- j) said-composition comprises poloxamer 188 (7% w/v) and polyethylene glycol 400 (2% w/v)
  - (i) wherein said-composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;

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k) said composition comprises poloxamer 188 (6% w/v), polyethylene glycol 400 (4% w/v), and propylene glycol (1% w/v):

- (i) wherein said-composition further comprises one or more compound selected from citric acid, disodium edetato, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal;
- I) said composition comprises poloxamer 188 (6% w/v), polyethylene glycol 400 (4% w/v), and propylene glycol (2% w/v):
  - (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbicidal; or

m) said-composition comprises poloxamer-188 (9% w/v) and polyethylene glycol 400 (2% w/v):

- (i) wherein said composition further comprises one or more compound selected from citric acid, disodium edetate, metabisulfate, benzyl alcohol, an antioxidant, a preservative, an antimicrobial agent, or a microbioldal.
- (Canceled)
- 8. (Previously Presented) The composition of Claim 1, wherein:
- a) said composition has a particle size diameter of between 25 and 200 nm;
- b) said composition has a particle size diameter of between 50 and 100 nm; or
- c) said composition forms particles of similar particle size.
- (Previously Presented) The composition of Claim 1, wherein:
- a) said composition does not support microbial growth;
- b) said composition is microbicidal; or

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c) said composition is sufficient for no more than a 10-fold increase in growth, of Staphylococcus aureus ATCC 6538, Escherichia coli ATCC 8739, Pseudomonas aeruginosa ATCC 9027 or Candida albicans ATCC 10231 for at least 24 hours.

- (Currently Amended) The composition of Claim 1, wherein:
- a) said composition is functionally <del>bioequilavent</del>bioequivalent to commercial lipid based anesthetic products:
  - (i) wherein said bioequilavencebioequivalence is demonstrated in dogs;
- (ii) wherein said <del>bioequilvence</del><u>bioequivalence</u> is demonstrated in humans; or
- b) said composition has a red blood cell blood plasma partition coefficient greater than that of commercial lipid based anesthetic products:
- (i) wherein said partition coefficient for said composition is between about 2 and 4.
  - 11. (Previously Presented) The composition of Claim 1, further comprising:
  - a) an acid;
  - b) a base;
  - c) a local anesthetic;
  - d) a second general anesthetic;
  - e) an antimicrobial agent;
  - f) a surfactant;
  - q) a tonicity modifier;
  - (i) wherein said tonicity modifier is glycerol;
  - h) a pH modifier; or
  - j) a second, third, fourth, fifth, or sixth excipient.
  - (Currently Amended) The composition of Claim 1, wherein said composition is substantially free of:

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a. a lipid, a long chain fatty acid, triacylglycerol, or glycerol ester;

b. a. an antimicrobial agent; or

e. b. a preservative.

13.-24. (Çanceled)

25. (New) A sterile aqueous pharmaceutical composition for parenteral administration of propofol, said composition comprising about 1% (w/v) propofol in combination with excipients, said excipients including:

7% to 9% (w/v) poloxamer component consisting essentially of Poloxamer 188; 2% to 4% (w/v) polyethylene glycol;

0% to 1% (w/v) propylene glycol; and

not more than 1% (w/v) lipid

wherein said excipients comprise not more than 15% (w/v) of said composition.

26. (New) The composition of Claim 25, wherein said excipients comprise: 8% (w/v) Poloxamer 188; 3% (w/v) polyethylene glycol 400; and 1% (w/v) propylene glycol.